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INDEX 'ADISALERTS, ADISINSIGHT, AGRICOLA, AIDSLINE, ANABSTR, AQUASCI,
BIGBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO,
CABA,
CANCERLIT, CAPLUS, CEABA, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU,
DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 12:39:40 ON 21 SEP
2000

SEA URIDINE(W)PHOSPHOGALACTOSE OR UDP-GALACTOSE

59 FILE AGRICOLA
1 FILE AIDSLINE
10 FILE ANABSTR
7 FILE AQUASCI
6 FILE BIGBUSINESS
1 FILE BIOCOMMERCE
1234 FILE BIOSIS
50 FILE BIOTECHABS
50 FILE BIOTECHDS
240 FILE BIOTECHNO
171 FILE CABA
91 FILE CANCERLIT
1819 FILE CAPLUS
6 FILE CEABA
1 FILE CEN
17 FILE CONFSCI
3 FILE CROPB
6 FILE CROPU
44 FILE DGENE
60 FILE DRUGB
11 FILE DRUGU
1 FILE EMBASE
167 FILE EMBASE
134 FILE ESPROBASE
1 FILE ESTI
16 FILE ECTA
267 FILE GENBANK
11 FILE IFIPAT
45 FILE JICST-EPLUS
219 FILE LIFESCI
640 FILE MEDLINE
2 FILE NTIS
3 FILE OSPAN
1 FILE PACT
512 FILE SCISEARCH
70 FILE TENLINE
240 FILE TMLIT
119 FILE USPATFULL
22 FILE WILDS
22 FILE WEINDEX

L1 QUE URIDINE W) PHOSPHOGALACTOSE OR UDP-GALACTOSE

FILE 'MEDLINE, EMBASE, SCISEARCH, BIOSIS, CAPLUS, TOXLIT, CANCERLIT'
ENTERED AT 12:41:47 ON 21 SEP 2000

L2 2713 S L1 AND SYNTHESIS OR BIOSYNTHESIS OR PROCESS OR PRODUCT?
L3 13 S L2 AND MORYNEBACTERIUM
L4 6 DUP REM L3 (9 DUPLICATES REMOVED)

LG ANSWER 1 OF 6 CAPLUS COPYRIGHT 2000 ACS

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000/019603	A2	20000525	WO 1999-US27599	19991118
W:	AE, AL, AM, AN, AO, AP, BA, BB, BG, BF, BY, CA, CH, CN, CR, CU, DE, DG, DK, DM, EE, EG, FI, GB, GD, GE, GH, GM, HP, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NA, NL, PL, PT, RO, RU, SE, SG, SI, SK, SL, TD, TM, TR, TT, TZ, UA, US, UZ, VN, YU, ZA, ZW, AM, AO, BY, EG, GE, GR, GU, HT, IL, IN, JP, KE, KG, KP, KR, KZ,			
RW:	GH, GM, KE, LS, MW, SD, SL, SN, TG, US, UZ, UA, AT, BE, CH, CY, DE, DK, ES, FI, FR, GR, GU, HT, IL, IN, JP, KE, KG, KP, KR, KZ,			
PRIORITY APPL. INFO.:	US 1998-109031 19981118 US 1998-109096 19981119			

This invention provides recombinant cells, reaction mixts., and methods for the enzymic **synthesis** of saccharides. The recombinant cells contain a heterologous gene that encodes a glycosyltransferase which catalyzes at least one step of the enzymic **synthesis**, as well as a system for generating a nucleotide sugar that can serve as a substrate for the glycosyltransferase. The nucleotide sugar may be supplied or synthesized by an enzymic pathway comprising a sugar nucleotide regeneration cycle. The reaction mixt. may contain a second cell type **producing** a nucleotide as a substrate for the sugar nucleotide regeneration cycle, preferably by a nucleotide synthase gene. Use of fusion proteins of glycosyltransferase and nucleotide sugar synthase combined with the use of an enzyme for substrate sugar **synthesis** is described. Chem. or enzymic sulfation may be used for the **synthesis** of sulfated sugars. The recombinant cells, reaction mixts., and methods are useful for efficiently synthesizing a large variety of saccharides, including polysaccharides, oligosaccharides, glycoproteins and glycolipids, using relatively low-cost starting materials. **Synthesis** of 3'-sialyllactose using *E. coli* expressing a CMP-sialic acid synthetase/ α .2,3-sialyltransferase fusion protein is described. Optional use of bakers yeast to **produce** CTP used in the sialic acid cycle and substrate for CMP-sialic acid synthase is also described. **Synthesis** of 3'-sialyllactose using *E. coli* expressing a CMP-sialic acid synthetase/ α .2,3-sialyltransferase fusion protein, GlcNAc 2'-epimerase, and sialic acid aldolase to synthesize CMP-sialic acid from GlcNAc is also described. Variations of the method using **Corynebacterium** expressing a CMP-sialic acid synthetase/ α .2,3-sialyltransferase

fusion protein and GTP-synthetase to **produce** the nucleotide, nucleotide sugar, catalyzing sugar transfer to acceptor saccharide is described. Finally, **synthesis** of trisaccharide Gal.alpha.1,3Gal.beta.1,4GlcNAc using **Corynebacterium** expressing UDP-glucose pyrophosphorylase, UDP-glucose-4'-epimerase, .beta.1,4-galactosyltransferase, and .alpha.1,3-galactosyltransferase is described.

L4 ANSWER 2 OF 6 SILENCE COPYRIGHT 2000 ISI (R)
ACCESSION NUMBER: 1999-49-481 MEDLINE
THE GENUINE ARTICLE: 443E
TITLE: Cloning and expression of beta 1,4-galactosyltransferase gene from *Helicobacter pylori*
AUTHOR: Endo T (Reprint); Kozumi S; Takata K; Ozaki A
CORPORATE SOURCE: KYOWA HAKKO KOGYO CO LTD, TOKYO RES LABS, 3-6-6 ASAHI KACHI, TOKYO 1448533, JAPAN (Reprint)
COUNTRY OF AUTHOR: JAPAN
SOURCE: GLYCOBIOLOGY, (AUG 2000) Vol. 10, No. 8, pp. 809-813. Publisher: OXFORD UNIV PRESS, GREAT CLAPENDON ST, OXFORD OX2 6DP, ENGLAND. ISSN: 0950-6858.
DOCUMENT TYPE: Article; Journal
FILE SEGMENT: LIFE
LANGUAGE: English
REFERENCE COUNT: 1

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

AB *Helicobacter pylori*, which is a human pathogen associated with gastric and duodenal ulcer, has been shown to express human oncofetal antigens Lewis X and Lewis Y. Although the mammalian glycosyltransferases that synthesize these structures are well characterized, little is known about the corresponding bacterial enzymes. We report that a novel beta 1,4-galactosyltransferase gene (HpgalT) involved in the **biosynthesis** of lipopolysaccharides in *H. pylori* has been cloned and expressed in *Escherichia coli*. The deduced amino acid sequence of the protein (HpGal-T) encoded by HpgalT consists of 274 residues with the calculated molecular mass of 31,751 Da, which does not show significant similarity to those of beta 1,4-galactosyltransferases from mammalian sources and *Neisseria*. It was confirmed that HpGal-T catalyzed the introduction of galactose from UDP-Gal in a beta 1,4 linkage to accepting N-acetylglucosamine (GlcNAc) residues by means of high-performance anion-exchange chromatography with pulsed amperometric detection (HPAEC-PAD). When the *E. coli* cells which overexpressed HpgalT was coupled with the UDP-Gal **production** system, which consisted of recombinant *E. coli* cells overexpressing its UDP-Gal **biosynthetic** genes and **Corynebacterium ammoniagenes**, N-acetylglucosamine, a core structure of lipopolysaccharide of *H. pylori*, was efficiently **produced** from ornithine, galactose, and GlcNAc.

L4 ANSWER 3 OF 6 MEDLINE
ACCESSION NUMBER: 1999-49-481 MEDLINE
DOCUMENT NUMBER: 443E-4081
TITLE: Large-scale **production** of N-acetylglucosamine through bacterial coupling.
AUTHOR: Endo T; Kozumi S; Takata K; Kakita S; Ozaki A
CORPORATE SOURCE: Tokyo Research Laboratories, Kyowa Hakko Kogyo Co., Ltd., Japan. enb.tetsuc@kyowa.co.jp
SOURCE: CARBOHYDRATE RESEARCH, (1999 Mar 31) 316 (1-4) 179-83. Journal code: CNY. ISSN: 0006-4215.
PUB. COUNTRY: Netherlands
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199-12
ENTRY WEEK: 19991202

AB A large-scale **production** system of N-acetylglucosamine, a core structure of various oligosaccharides, was established by a whole-cell reaction through the combination of recombinant *Escherichia coli* strains and *Corynebacterium ammoniagenes*. Two recombinant *E. coli* strain over-expressed the UDP-Gal **biosynthetic** genes and the beta-(1->24)-galactosyltransferase gene of *Neisseria gonorrhoeae*, respectively. *C. ammoniagenes* contributed the **production** of UTP from orotic acid. N-Acetylglucosamine was accumulated at 279 mM (107 g L⁻¹) after a 38 h reaction (2.5 L in volume) starting from orotic acid, D-galactose, and 2-acetanido-2-deoxy-D-glucose.

L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2000 ACS

ACCESSION NUMBER: 1998:197631 CAPLUS

DOCUMENT NUMBER: 129:056412

TITLE: **Processes for producing sugar nucleotides and complex carbohydrates**

INVENTOR(S): Koizumi, Satoshi; Sasaki, Katsutoshi; Endo, Tetsuo; Tabata, Kazuhiko; Ozaki, Akio

PATENT ASSIGNEE(S): Kyowa Hakko Kogyo Co., Ltd., Japan; Koizumi, Satoshi; Sasaki, Katsutoshi; Endo, Tetsuo; Tabata, Kazuhiko; Ozaki, Akio

SOURCE: ECT Int. Appl., 119 pp.

CODEN: PIXXDL

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9611343	A1	19960316	WO 1997-JP3226	19970312
W: AU, BG, BR, CA, CN, CZ, DE, HU, JP, KE, MX, NO, NZ, PL, RO, SG, SI, SK, UA, US, VN, AM, AQ, BY, EG, KE, MD, RU, TJ, TM				
EW: AT, BE, CH, EE, FR, ES, FI, FR, GP, GR, IE, IT, LI, MC, NL, PT, SE				
CA 2037849	AA	19960401	CA 1997-1137649	19970312
AU 9742263	A1	19960414	AP 1997-41275	19970312
EP 870841	A1	19961014	EP 1997-41036	19970312
E: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
CN 1017135	A	19990103	CN 1997-191696	19970312
PRIORITY APPL. INFO.:				
			JP 1996-144451	19960317
			JP 1996-185666	19961028
			WO 1997-JP3226	19970312

AB Sugar nucleotides are manufd. with microorganism or enzyme **producing** NTP from nucleotide precursor and with microorganism or enzyme **producing** sugar nucleotides from sugar and NTP. Complex carbohydrates are manufd. with the described microorganism/enzyme and microorganism/enzyme that **produces** complex carbohydrates from sugar nucleotide and complex carbohydrate precursor. Also given was **prodn.** of N-acetylglucosamine-1-phosphate with galactokinase-high microorganism.

L4 ANSWER 1 OF 6 MEDLINE

DUPLICATE 1

ACCESSION NUMBER: 1993414050 MEDLINE

DOCUMENT NUMBER: 96414050

TITLE: Large-scale **production** of UDP-**galactose** and globotriose by coupling metabolically engineered bacteria.

AUTHOR: Koizumi S; Endo T; Tabata K; Ozaki A

CORPORATE SOURCE: Tokyo Research Laboratories, Kyowa Hakko Kogyo Co., Ltd., Machida, Japan.. skoizumi@kyowa.co.jp

SOURCE: NATURE BIOTECHNOLOGY, (1996 Sep; 14 (9) 847-50. Journal code: CQ3. ISSN: 1087-0156.

PUB. COUNTRY: United States

(Journal; Article; JOURNAL ARTICLE)

LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199901
ENTRY WEEK: 19990104

AB A large-scale **production** system of uridine 5'-diphospho-galactose (UDP-Gal) has been established by the combination of recombinant

Escherichia coli and *Corynebacterium ammoniagenes*. Recombinant *E. coli* that overexpress the UDP-Gal **biosynthetic** genes *galT*, *galK*, and *galU* were generated. *C. ammoniagenes* contribute the **production** of uridine triphosphate (UTP), a substrate for UDP-Gal **biosynthesis**, from orotic acid, an inexpensive precursor of UTP. UDP-Gal accumulated to 72 mM (44 g/L) after a 21 h reaction starting with orotic acid and galactose. When *E. coli* cells that expressed the α gal,4-galactosyltransferase gene of *Neisseria gonorrhoeae* were coupled with this UDP-Gal **production** system, 372 mM (188 g/L) globotriose (Gal α 1gal-4Gal β 1gal-4Glc), a trisaccharide portion of verotoxin receptor, was **produced** after a 35 h reaction starting with orotic acid, galactose, and lactose. No oligosaccharide by-**products** were observed in the reaction mixture. The **production** of globotriose was several times higher than that of UDP-Gal. The strategy of **producing** sugar nucleotides by combining metabolically engineered recombinant *E. coli* with a nucleoside 5'-triphosphate **producing** microorganism, and the concept of **producing** oligosaccharides by coupling sugar nucleotide **production** systems with glycosyltransferases, can be applied to the manufacture of other sugar nucleotides and oligosaccharides.

L4 ANSWER 6 OF 6 MELLINE DUPLICATE 1

ACCESSION NUMBER: 9718090A MELLINE

DOCUMENT NUMBER: 9718090P

TITLE: The *galE* gene encoding the UDP-galactose 4-epimerase of *Brevibacterium lactofermentum* is coupled transcriptionally to the *dmdF* gene.

AUTHOR: Quirica J A; Marcos A T; Malumbres M; Martin J F

CORPORATE SOURCE: Department of Ecology, Genetics and Microbiology, Faculty of Biology, University of Leon, Spain.

SOURCE: GENE, (1998 Oct 14) 177 (1-2) 103-7.

Journal code: EMB. ISSN: 0378-1119.

PUB. COUNTRY: Netherlands

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

OTHER SOURCE: GENBANK-242413

ENTRY MONTH: 199701

AB The *galE* gene of *Brevibacterium lactofermentum*, encoding UDP-galactose 4-epimerase (EC 5.1.3.1), has been identified by DNA sequencing downstream from the *orf1-sigB-dmdF* region. The arrangement of the *sigB-dtxE-galE* cluster is also conserved in *Corynebacterium diphtheriae*. The deduced *galE* **product** was a protein of 329 aa residues (35.4 kDa) that shared a high degree of identity to known UDP-galactose 4-epimerase proteins from Gram-positive microorganisms (*Streptomyces lividans* and *Streptococcus thermophilus*). Transcriptional analysis of the *dmdF* and *galE* genes in nutrient-rich medium showed that these genes are part of an operon, that is actively transcribed as a bicistronic mRNA during the exponential growth phase,

but

transcription of the operon is decreased during the stationary growth phase. In addition, the *dmdF* gene was also expressed as a monocistronic 0.7-kb transcript during the active growth phase.

=> d his

L11 ANSWER 37 OF 42 TOXLIT
ACCESSION NUMBER: 1990:98326 TOXLIT
DOCUMENT NUMBER: CA-113-206233P
TITLE: Cloning and expression of **cDNA** for human
membrane-bound beta-1,4-**galactosyltransferase**.
AUTHOR: Fukuda MN; Appert HA
SOURCE: (1990). PCT Int. Appl. PATENT NO. 90 07000 06/28/90 (La
Jolla Cancer Research Foundation).
FUB. COUNTRY: United States
DOCUMENT TYPE: Patent
FILE SEGMENT: CA
LANGUAGE: English
OTHER SOURCE: CA 113:206233
ENTRY MONTH: 199012
AB A full-length **cDNA** encoding the membrane-bound form of beta-1,4-
galactosyltransferase from human Golgi bodies is cloned and
expressed in *Escherichia coli* and antibodies raised to peptides
from the protein. The enzyme is involved in post-translational
modification of proteins and there are pathol. consequences from
deficiencies in the enzyme (congenital dyserythropoietic anemia type II).
The full-length **cDNA** was constructed from a pair of overlapping
clones from a human placental **cDNA** library in *lambdagt11* and
expressed in *E. coli* using pIN-III-ompA3 as the expression
vector. Antibodies to a peptide from the carboxy-terminal region of the
protein were raised in rabbits by conventional methods.

L11 ANSWER 39 OF 42 CAPLUS COPYRIGHT 2000 ACS

ACCESSION NUMBER: 1991:1540 CAPLUS

DOCUMENT NUMBER: 114:1540

TITLE: Sequence of a **cdna** encoding human
galactose-1-phosphate uridyl transferase

AUTHOR(S): Flach, James E.; Reichardt, Juergen K. V.; Elsas,
Louis J., II

CORPORATE SOURCE: Dep. Pediatr., Emory Univ., Atlanta, GA, 30322, USA

SOURCE: Mol. Biol. Med. (1990), 7(4), 365-9

CODEN: MBIMDG; ISSN: 0735-1313

DOCUMENT TYPE: Journal

LANGUAGE: English

AB A revised sequence of a **cdna** that encodes a human
galactose-1-phosphate uridyl transferase is reported here. The
cdna is 1295 bases in length and encodes a 43,000 Mr protein. The
sequence was derived from a **cdna** clone isolated from a
transformed human lymphoblast cell line and amplified in a polymerase
chain reaction. The revised sequence reveals a higher degree of amino
acid conservation between the human enzyme and the homologous enzymes

from
Escherichia **coli** and yeast than was previously thought to exist.

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Search Results -

Terms	Documents
L11 and Corynebacterium	2

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Derwent World Patents Index

Database: IBM Technical Disclosure Bulletins

L11 and Corynebacterium

Refine Search:

Clear

Search History

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<u>DB Name</u>	<u>Query</u>	<u>Hit Count</u>	<u>Set Name</u>
USPT,JPAB,EPAB,DWPI	L11 and Corynebacterium	2	L13
USPT,JPAB,EPAB,DWPI	L11 and Corynebacterium	2	L12
USPT,JPAB,EPAB,DWPI	L4 and galactose	87	L11
USPT	4296203.pn.	1	L10
USPT	5516665.pn.	1	L9
USPT	5409817.pn.	1	L8
USPT,JPAB,EPAB,DWPI	L5 and orotic acid	0	L7
USPT,JPAB,EPAB,DWPI	L2 and orotic acid	1	L6
USPT,JPAB,EPAB,DWPI	L2 and galactose	67	L5
USPT,JPAB,EPAB,DWPI	L1 and orotic adj acid	378	L4
USPT,JPAB,EPAB,DWPI	L1 and orotic adj acid	378	L3
USPT,JPAB,EPAB,DWPI	L1 and (sugar adj nucleotide)	185	L2
USPT,JPAB,EPAB,DWPI	synthesis OR biosynthesis OR product? Or process? same (sugar adj nucleotide)	925064	L1

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Search Results -

Terms	Documents
Corynebacterium and l3	4

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Corynebacterium and l3

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USPT,JPAB,EPAB,DWPI	Corynebacterium and l3	4	L6
USPT,JPAB,EPAB,DWPI	synthesis OR biosynthesis same l3	252198	L5
USPT,JPAB,EPAB,DWPI	(Prepara? OR Process or making or manufacture) same l3	7	L4
USPT,JPAB,EPAB,DWPI	UDP-galactose	121	L3
USPT,JPAB,EPAB,DWPI	phosphogalactose	4	L2
USPT,JPAB,EPAB,DWPI	(uridine adj phosphogalactose) OR (Uridine adj phosphoglucose)	0	L1